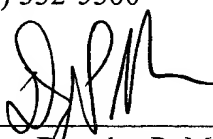


If a telephone conference would be helpful in resolving any issues concerning this communication, please contact Applicants' primary attorney-of record, Douglas P. Mueller (Reg. No. 30,300), at (612) 371.5237.

Respectfully submitted,

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DPM/rw



WE CLAIM:

1. A method of analyzing argentinated peptides or proteins using mass spectrometry comprising:
  - (a) combining an oligopeptide with silver to provide a sample comprising argentiated oligopeptide;
  - (b) submitting the sample to a mass spectrometer;
  - (c) performing scans of silver containing peaks in optimum collision energies;
  - (d) identifying any doublet or triplet peak pattern;
  - (e) confirming with Y ions;
  - (f) determining partial sequence by the mass separation between two successive doublet or triplet pattern.
2. A method according to claim 1 wherein the performing scans comprises collecting product ion spectra of the  $[M + Ag]^+$  ion, where M = oligopeptide;
3. A method according to claim 1 [or 2] wherein the oligopeptide comprises from about 3 to about 10 amino acids.
4. A method according to [any one of claims 1-3] wherein the silver is silver nitrate. CLAIM 1 --
5. A method according to [anyone of claims 1-4] wherein the determination of partial sequence comprises searching for, and identifying cleaved amino acid residues based on differences in  $m/z$  values of neighboring triplets where the  $m/z$  value of the  $[b_n - H + Ag]^+$  ion and the corresponding  $[y_n + H + Ag]^+$  ion are related by the formula:  $[y_n + H + Ag]^+ = [M + Ag]^+ + Ag^+ - [b_n - H + Ag]^+$ . CLAIM 1 --
6. A method according to claim 5 wherein the searching and identifying is conducted by a custom search algorithm.
7. A method according to claim 6 wherein the algorithm is written in

Visual Basic and looks for the triplet peak pattern of  $(m/z)_1$ ,  $(m/z)_2 = (m/z)_1 - 18.0$ , and  $(m/z)_3 = (m/z)_2 - 28.0$  as well as the doublet pattern of  $(m/z)_2$  and  $(m/z)_3$ , all to within  $\pm 0.5 m/z$  unit.

5 8. A method according to anyone of claims 1-7 wherein product ion spectra of the  $[M + Ag]^+$  ion are collected under  $E_{cmS}$ , of 1.5, 2.0, 2.5 and 3.0 eV.

10 9. A method according to anyone of claims 1-8 wherein the mass spectrometer is a triple quadrupole mass spectrometer, two triple quadrupole mass spectrometers, a quadrupole/time-of-flight mass spectrometer, an ion-trap mass spectrometer, or a time-of-flight mass spectrometer amenable to post-source decay or collision-induced dissociation.

10. A method of analyzing argentinated peptides or proteins using mass spectrometry comprising:

- 15 (a) combining an oligopeptide with silver nitrate in solution;  
 (b) submitting a sample of the solution to a mass spectrometer;  
 (c) collecting product ion spectra of the  $[M + Ag]^+$  ion, where M = oligopeptide;  
 (d) identifying a triplet peak pattern;  
 20 (e) identifying a doublet peak pattern;  
 (f) searching for, and identifying cleaved amino acid residues based on differences in  $m/z$  values of neighboring triplets where the  $m/z$  value of the  $[b_n - H + Ag]^+$  ion and the corresponding  $[y_n + H + Ag]^+$  ion are related by the formula:  $[y_n + H + Ag]^+ = [M + Ag]^+ + Ag^+ - [b_n - H + Ag]^+$ .  
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11. A method according to claim 10 wherein the oligopeptide comprises from about 3 to about 10 amino acids.

12. A method according to claim 10 or 11 wherein the searching and identifying is conducted by a custom search algorithm.

13. A method according to claim 12 wherein the algorithm is written in Visual Basic and looks for the triplet peak pattern of  $(m/z)_1$ ,  $(m/z)_2 = (m/z)_1 - 18.0$ , and  $(m/z)_3 = (m/z)_2 - 28.0$  as well as the doublet pattern of  $(m/z)_2$  and  $(m/z)_3$ , all to within  $\pm 0.5 m/z$  unit.

5 14. A method according to anyone of claims 10 to 13 wherein product ion spectra of the  $[M + Ag]^+$  ion are collected under  $E_{cmS}$ , of 1.5, 2.0, 2.5 and 3.0 eV.

10 15. A method according to anyone of claims 10-14 wherein the mass spectrometer is a triple quadrupole mass spectrometer, two triple quadrupole mass spectrometers, a quadrupole/time-of-flight mass spectrometer, an ion-trap mass spectrometer, or a time-of-flight mass spectrometer amenable to post-source decay or collision-induced dissociation.